

U9/341550 PCT/US98/00840

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224 10	GCG	GCC ACC ACC CAG A T T Q	, CC 7	ACC		CCG P	CCG GCC A	AGC S	ACC T	SCG P	CTG	GCG A	AAC	TCG	TCG	CTG L	TAC Y	GTC V	GGT G	GAC D	283 29	
284 30	CTG L	GAG A E K	AAG ( K	GAT D	GTC V	ACC	GAG	A GCC	cAG Q	CTG	TTC	GAG	CTC	TTC	TCC	TCG	GTT V	0 0 0	CCT P	GTG V	343 49	
344 50	GCC	TCC A S I	ATT ( I	) G	GTG V	TGC C	7 7 7	GAT D	GCC A	GTC	ACG T	ACG CGC T R	CGC R	CGC TCG R	CTG	0 0	TAC Y	TAC GCC Y	ľAC ľ	GTC V	403 69	
<b>1</b> 04 70	AAC N	TAC A Y N	AAC N	AGC	GCT A	CTG L	GAC	CCC	cAG Q	GCT A	GCT	GAC	CGC	GCC A	ATG M	GAG	ACC	CTG L	AAC N	TAC Y	463 89	
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CCG P	TAC Y	ATG M	9 9	CAG Q	GCG A	TCC S	CCG P	ATG M	GAC D	AGC	
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CCG P	CCC	CCG P	ATG M	CCC P	GCT A	GCG A	GCG A	CAG Q	CTG L	CTC	
CCG P	ည ဗ	CCG P	ATG M	ට ව	GCC A	GCG A	9 <b>A</b>	CTG	CIT	GTG V	
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2159 8		2284 43	2364	2444	2516 8	2576 28	2636 48	2715 49	2790 5	2846 23	
2089 GGCTGGTGGCGCGCGCGCGCGCTGCTTGGGCCGCCGCGCGC ATG GGC GCG GCG GCG GTG TGG  1 1 2089 GGCTGGTGGCGCGCGCGCGCGCGCGCGCCGCCGCCGCCAGC ATG GCG GCG GCG GTG TGG  1 2089 GGCTGGTGGCGCGCGCGCGCGCGCGCGCCGCCGCCGCCAGC ATG GCG GCG GCG GTG TGG	$2160$ gag cag tige ittig ctig ctit ctig gee gee gige aag ceg ege cga actig gee gae gee agg $9~{ m E}$ Q C L L L L A A V K P R R T G A D G R	2220 CTG GCG TTG ACG CCG CGC CAC AAC ACA AAG TTG GTG GCG TGA AAGTCTCTGGGCGTGCTCCG $29~\mathrm{L}$ A $^\star$	2285 GACGGTTGTAAGGTTTTTAAGAACTGGCTTTTTGGCCGGGTTGCCGCCCCAAAGGCGGAACGGCGGTCTTTTCAGGCCAATCA 2364	2365 CATCCGGCTGGAAAATTCTTACCAAAGCCAACCCTGCACCCAAAAATTTCGGGTTCCGAAAGAACACTCCCCTTTTTT	2445 CCGCCAACGCGTTCTTTCAAGGCCAATCACTTTCCGGGTTGGAAGAAA ATG TTA CCC GGA AAA GGC GGG AAG	2517 CCC CCT GCA CCA GTT ATT CGG GGT TTC GCC GGG AAT GAG CAA GCG TTC GGG CTG	2577 TYG GCC GTA TCG CGA ACG CTG TCG GGG TGT CAG GCG CCA GAA GGA AGG ATG ACG TTT TGG 29 L A V S R T L S G C Q A P · E G R M T F W	2637 TGA AGGGGTGCAAACTGAGCACACGAGTTTTGGCAATAGACGTGGAAAAGTCCAGTGCGGGGTGAGGCGGATAGCGGA 2715 49 *	2716 ATCAAGCGTGCGGGTCCCTGGCGAGACGATTCTTTTTTTT	2791 TGT TTT GAG CAG ACT GTA AAG TGC CCG ACG CTA AAA AAG CGG CCG CGA ATT CC 6 C F E Q A T V K C P T L K K R P R I	

ggagtccctg tggcgagctg

ccacccagga ggttcgttga

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KDDDVDVTVVTVKNWDETVKKSKFALVEFYAPWCGHCKTLKPEYAKAATALKAAAPDA

LIAKVDATQEESLAQKFGVQGYPTLKWFVDGELASDYNGPRDADGIVGWVKKTGPPA

VTVEDADKLKSLEADAEVVVVGYFKALEGEIYDTFKSYAAKTEDVVFVQTTSADVAKA

AGLDAVDTVSVVKNFAGEDRATAVLATDIDTDSLTAFVKSEKMPPTIEFNQKNSDKIF

NSGINKQLILWTTADDLKADAEIMTVFREASKKFKGQLVFVTVNNEGDGADPVTNFFG

LKGATSPVLLGFFMEKNKKFRMEGEFTADNVAKFAESVVDGTAQAVLKSEAI PEDPYE

DGVYKIVGKTVESVVLDETKDVLLEVYAPWCGHCKKLEPIYKKLAKRFKKVDSVIIAK

MDGTENEHPEI EVKGFPTILFYPAGSDRTPIVFEGGDRSLKSLTKFIKTNAKI PYELP KKGSDGDEGTSDDKDKPASDKDEL

gctgctgctg tgtggtgacc gttctacgct caccgccctg gagtacgttt acgccatgaa ccgttggaac cttcttgccc ttaccctggg gctcatgctt ccgatgagta gacgacgacg tcgacgttac cgcttgtgga ctaaggctgc gtcaagaact gggatgagac cgtcaagaag tccaagttcg gacceteaag cetgagtaeg gtggcagcgc ccttcaccaa gcaccagttt gaggaggacg atgccccgc cgccctaag gccactgcaa ccttggtgcg 181 241

agaccaccag cgccgacgtc gaagaagaag ggaggcggac gegagateta egaeaeette ttggctgggt tgaagteeet gtgttcgtgc gccgacaagc gatggcattg gccctggagg ccgcgacgct ctacttcaag aagteetaeg eegeeaagae egaggaegtg actggcccc ccgccgtgac cgttgaggac gggctacccc gctgaggtcg ttgtcgtcgg gcccagaagt tcggcgtgca gettetgact acaaeggeee 361 541 421 481

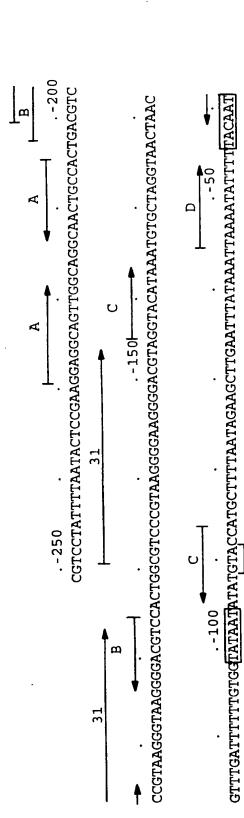
FIG. 2A

RECTIFIED SHEET (RULE 91)

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2B FIG. gctgcaagaa agatgaacaa gatagcgccg ggatcaggga gtaagcggct gcttggagca ggctaggggc tgcgactgga tggatggatt ggatgtgcgt cgcccggcac ctacaagaag tggcactgag cctgaccaag cgacggcgac cactgtgcat cggtgcagtc ggacgtgctg tcccgccggc cttcgccggt gaccgcgttc cctgaaggcc ccagctggtg gaagttccgc ggtggacggc ggatggcgtc tgacaagatc cttcggcctc ccgcagcgca tgcgcttgca gcgggcgcat tccacagccc tgtgtgcata ctgtcgccct tcgcggcgtg caaagaacgg acccctatga acgagaccaa ccaagatgga tcctgttcta cgctcaagtc agaagggctc cgggaggaag cctctccctg gtgtatgtgg ttgtcgggtc tcaccaactt agaagaacaa ccgagagcgt tggagcccat aggacgagct tcggttggtg ccgccgacga agttcaaggg tggtcaagaa agaagaactc ctgactccct ttccctacca ggcgaccgct ttggtacggt ctcggcgctt agccgggcct cagtgtgcgg gtggttctgg gagctgccca cttgcgcgct gaggctgcgc ctgacggtgt agcgagggtg gcgagggcaa gccagcaaga gccgaccccg tgcaagaagc gtcatcatcg gcgtccgaca caacggcgga tgctgccggg ggcgcccgtg agggacgctg ttcttcatgg gctaagttcg atccccgagg accgtgtccg gacatcgaca gagttcaacc ctgtggacca ggagaggata gctgtgttgc gggattgcga gcgaggaaaa ggtggattcc ctggcatgcg ggtagtggtg tactagtatg tctgttgagg ggatgatgag ggtcaagggc gatecegtae ggacaagccc ggagcgtctg acggagcacg tttgccaaag gtgcggccac gttcgagggc ctggcagcag gcagctgatt gggcgacggc gtcggaggcc cgtggagtct cgccgtggac cccaccatt gttccgcgag gctgctgggc tgacaacgtg cctggccacg gctggagtca agctagcgca gacgcacggt cggagtgcat ggctgttgct gccctgcggt catgaagagt tgtccggatg ccgagatcga ccaacgccaa cggacgacaa ccccaggttt gtgatccgtc gcgttacggg ccggcctgga ccaccgccgt tcatgactgt cctcgcctgt ccgtgctcaa tgggcaagac acgcccctg gctttaagaa ccccatcgt taaggaggag aagacgagac agaagatgcc gcatcaacaa tcaacaacga agttcacggc ggccgcgtga cagcggatcg ggagccaagg ctggccaagc agcgaccgca agttttttag cgtttctctc gtgccccgac ggttccgaac cgccttgcgc gacgccgaga aagggcgcca atggagggcg tacaagattg ctggaggtgt aacgagcacc ttcatcaaga gagggcacct atctgaacta ggatgggagt ccggcagcgc gctggcgagc agagatgaga cttgctagga ttcgtgaccg accgcgcagg gccaaggccg gaggaccgcg gtcaagtcgg ttcaacagcg 2401 561 101 161 221 441 501 801 861 981 041 2281 901 261 381 621 681 741 841 961 201 321 921 081 141

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10 Thr Ala Ile Leu Glu Arg Arg Glu Asn Ser Met

TCT AGC CTA TGG GCT CGT TTT TGT GAG TGG ATC ACT TCA ACT GAA AAC CGT TTA TAC ATC GGT TGG TTC GGT GTA ATC ATG ATC CCA TGT CTT CTT ACT GCA ACA TCA GTA TTC ATC ATC Leu Tyr Glu Asn Arg Thr Glu Trp Ile Thr Ser Asn Ser Ser Leu Trp Ala Arg Phe Cys . 50 G1y100

50 Ser GCT TTC ATC GCT GCT CCG CCA GTA GAC ATC GGT ATC CGT GAA CCA GTT TCA GGT Glu Pro Val Ser Gly Ser Val Phe . -200 Ala Thr Pro Pro Val Asp Ile Asp Gly Ile Arg Cys Leu Leu Thr Pro ( Ile Met Ile Leu Ile Ala Ala Gly Val Phe Trp Gly

FIG. 3A

230

GGT

GCA

TCT AAC

CCA ACT

GGT GCT GTA ATC

GGT AAC AAC ATC ATT ACA

CTT TAC

Len

Len

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110 90 Gly GGT Try Asn Gly GGT TAC AAC Asn Ser Asp Glu Trp Leu TGG TTA Pro Thr CTA GAC GAG Ile Val Leu IleVal Ala Ser TGG GAA GCT GCT TCT .300  $_{
m G1y}$ Ala Ala Ile Thr Ser Trp Glu Ile I1eTAC CCA ATT Asn Pro Asn Tyr G1yPhe TyrCAC Leu

130 .450 Arg Glu GTA TAC TGC TAC ATG GGT GGT GAG G1X]/ Met Tyr Cys TyrAla Val GGTG1yCTA Leu Leu TIC CIT Cys His Phe TGT CAC Leu GTT Val Leu Ile CIT ATC Gln CAA TAC Tyr Pro CCT

150 Pro GTA GCT TAC TCA GCT CCA Ser Tyr Ala Val GCT Ala ATC Ile TTA GGT ATG CGT CCA TGG Pro Trp Arg Met GlyLeu TCT TTC CGT Ser Phe Arg TTA Leu GAA Glu ŢĞĞ Trp

170 Asp CAC GAC GAA Ser TCA TTC Ser Phe ATC GTA TTC CAA GCA Gln GGC CAA GGT Gln Gly Tyr Pro Ile Gly TAC CCT ATC ccr rra ger][Arc rcr ggr ACr rrc AAC rrc ATG GTTVal Ile Phe Leu TTC TTA GTA Ser Ala Val TCA GCT Thr GCT Ala GCA Ala GTA GCT Ala Val

190

His

Glu

Ala

Phe

Ile Val

Met

Phe

Asn

Phe

Thr

 $_{
m Gly}$ 

Ser

Ile

Gly

Len

Pro

Gly

210 TTA Ser GGT GTA TTC GGT GGT TCA  $_{
m Gly}$ G1yPhe Val Gly Ala TTA GGT GTT GCT Leu Gly Val .600 CTT ATG CAC CCA TTC CAC ATG Met His Phe Pro His Len AAC ATC Ile Asn

GAA AAC Thr ACT ACA Thr GAA Glu Arg GGT TCT TTA GTT ACT TCA TCT TTA ATC CGT Ile Ser Leu Ser Thr Ser Leu Val Gly ATG CAC His Met Ala TCA

Ala 250 GCT Val GTA Tyr Asn Ile TAC AAC ATT Thr ACT GAA Glu Glu Glu GAA GAA CAA Gln GGT Gly TTC Phe CGI Tyr Arg TAC GGTSer Ala Asn Glu Gly AAC GAA .700 GCT TCA

Glu

GAA

.750

TCA CGT AAC TCT TAC GCT TCT TTC AAC TTC CAA CGT CTA ATC gcr car][ggr rac rrr ggr

Ser Arg Asn Ser Asn Phe .850 Ser Ala Tyr Gln Phe Ile Len Arg Gly Phe Tyr His Gly

290 Leu TTA GGT  $_{
m Gly}$ TTA Ala Leu GGT ATT TGG TTC ACT GCT Trp Phe Thr Ile G1yIle TTA GCT GCT TGG CCG GTA ATC Val Pro Val Trp Ala Ala Leu TIC Phe TIC Phe

CAC

Ala

His

Leu

TCA

Ser

.900

310 Gln CAA Ser CAA TCA GTA GTA GAC TCA Val Asp Gln Ser Val GGT TTC AAC TTC AAC Asn Phe Gly Phe Asn TTC AAC TTA AAC Asn Leu Asn Phe GCA Ala Met ACT  $\operatorname{Thr}$ 

330 Val GTA GAA CTA AAC ACT TGG GCA GAC ATC ATC AAC CGT GCT AAC TTA GGT ATG Met GlyLeu Asn Ala Arg Asn Ile Ile Trp Ala Asp .950 Thr Leu Asn CGT GTA Val Arg ggT

.1050 Ser 350 TCA AGC Ser CGT AAC GCT CAC AAC TTC CCT CTA GAC TTA GCT TCA ACT AAC TCT Asn Ser Pro Leu Asp Leu Ala Ser Thr His Asn Phe Ala Arg Asn .1000 GAG Glu His ATG CAC

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Ala

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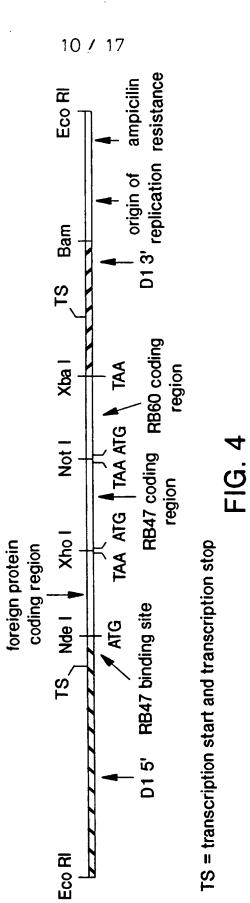
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Asn

Asn

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60 20	120 40	180 60	240 80	300	360 120	420 140	480 160	540 180	600	660
CGT R	CCG P	CTG	GTC V	GCT A	ATC	AAG K	AAG K	GTG V	ATT I	AGG R
GGT	ACC	CAG Q	GCC A	CAG Q	CGC R	ATC I	ეტ ე	TTC	AAG K	GCA A
GAA E	AGC S	GCC A	GAT D	CCC	ATG M	TTC F	TTC	ည ဗ	AAG K	AGG R
ATC I	GCC A	GAG E	CGC R	GAC D	CCT P	ATC I	GCC A	TAC Y	CAG Q	CCC
CAT H	CCG	ACC	TGC C	CTG L	AAG K	AAC N	TCG	) 9	AAC N	CGC R
0 0 0	CAG Q	GTC V	GTG V	GCT A	ე ე	ည ၅	TTC	AAG K	GTC V	GAC D
AGC S	ACC	GAT D	CGC R	AGC S	AAC N	GTC V	ACC T	TCG	ACC T	GCT A
AGC S	ACC T	AAG K	ATT I	AAC N	GTG V	0 0	GAC D	GTG V	CAG Q	CGC R
CAC H	GCC A	GAG E	TCC S	TAC Y	GTC V	TCG	CAC H	ეტე ე	ATT I	AAG K
CAT H	GCG A	CTG L	GCC	AAC N	CAT H	AAG K	CTG L	AAC N	GCC A	CAG
CAT H	CCG P	GAC	GTG V	GTC V	TAC Y	CGC	GCC A	GCC A	2 2 3 3	TTC
CAT H	BCC A	GGT G	CCT P	TAC Y	AAC N	GCC A	AAG K	GAC D	GAT D	CCC
CAT H	TCG S	GTC V	ეე ე	GCC A	CTG L	TCG	GCC A	ACT T	GCC A	GCC A
CAT H	JCC S	TAC Y	GTT V	TAC Y	ACC T	CCT P	GAC D	GCC A	GCT A	GTG V
CAT H	GAG E	CTG L	TCG	ე ე	GAG E	GAC D	ATC I	GTT V	GCC	TAC Y
CAT H	ACT	TCG S	TCC S	CTG L	ATG M	0 20 20	ACC	AAG K	CAG Q	GTG V
CAT H	ACT	TCG	TTC F	TCG	GCC	CAC H	AAG K	73GC C	GAC D	ATC I
CAT H	GCG A	AAC N	CTC L	CGC R	CGC R	TCG S	GAC D	TCC	GAG E	AAG K
ည ဗ	ATG M	906 <b>A</b>	GAG E	CGC R	GAC D	TGG W	CTG L	CTG L	TTC	ე ე
1 ATG 1 M	CAT H	CTG L	$\operatorname{TTC}_{\mathbf{F}}$	ACG	GCT A	ATG M	AAC N	ATT I	CAC H	GAG E
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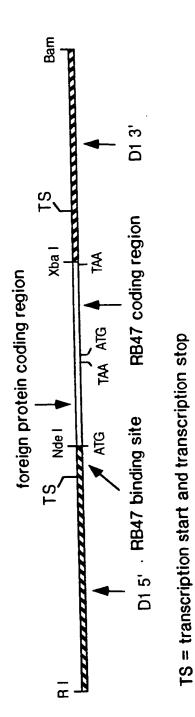
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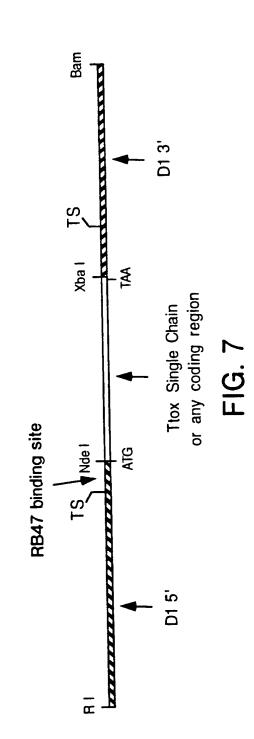
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720 240	780 260	840 280	300	960 320	1020 340	1080 360	1140 380	1200 400	1260 420	1278 424
CTG L	AAG K	TGC	CAG Q	CGT R	GAC	ATG M	GAG E	TAC Y	CAG Q	
GAG E	GAC D	AAG K	GCC A	GAG E	GTC V	GTC V	GAC D	CTG L	ATG M	
GAC D	GAC	GCC	CGC R	CAG Q	GAG E	AAG K	CCAC H	CCC	AAC	
GAC	AAG K	GCG A	ე ე	AAG K	GAG E	TGC C	AGC	AAG K	GCC A	
ე ე	ATG M	TCG	GCC	AGC S	GAC D	TCG	ACC T	0 0 0	GAG	
ATC I	GTC V	GAG E	TAC Y	GAG E	TCC	ACC	TTC	AAG	CTG	
GAC D	GTG V	GCC A	CTG L	GAG	CTG L	ATC	13GC C	GTC	CAG	
GCC A	GCG A	GAC D	ACC	GCC	AAC N	ACC	GTG V	ATG M	ACC T	
CCG P	AGC S	AAG K	AAG K	AAG K	AAG K	ට වි -	TTC	AAG K	, GCC	5B
TTG	ACC	TTC	ე ე	CAG Q	GTC V	TCT	ე ე ე	ე ე	CGT R	FIG.
AAC N	ATC I	AAC N	AGC S	CGC	TAC	AAC	TTC	AAC N	S CGC	ш
AAG K	GAG E	ATC I	ATG	CTG	CTG	GCC A	ეენ ენენ 1	, ATG M	; GTG V	
GTC V	ე ე	TTC	GAG	ATG M	AAC N	TTC F	: AAG K	GAG E	GAC D	
TTC	CAC	ည တ	CGC R	GCG A	ATG M	; cris	) TCC S	, ACC T	AAG K	
GTG V	GAG	TTC	GAG	GAG	AGC S	GAG E	AAG K	GTG V	CGC R	GGATCC
AAC N	ACC	ეტე ე	AAC N	r cgc	CAG	; cgt R	2 9 9	S CCC	cAG Q	1 GG
ACC	GCC A	AAG K	CTG	GAG	TAC Y	CTG	AGC S	CCG	GCG A	TAA *
TAC	ATG M	AGC	TAC	ACC	AAG K	GCC A	၁၅၅	CGG R	CTG	: ATG M
TTG	AAG K	ည ဗ	GAG	AAG K	CIG	GAC	GAC D	ACC T	GCC A	CGC R
ACG T	ည္	0 0 0	GTG	AAG K	TAC Y	GAC D	AAG . K	. GCC	GTG V	GCG
661 221	721 241	781 261	841 281	901 301	961 321	1021 341	1081 361	1141 381	1201 401	1261 421

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Garage Contract Contract

FIG. 6





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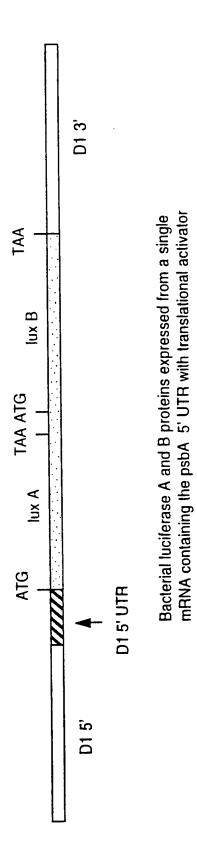
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C.	reinhardt	ii expre	ssea	
3 soluble	3 pellet	2.1 soluble	2.1 pellet	et Tox Fab
		*****	<b></b>	

Fab

Self-are committee

FIG. 8



element.

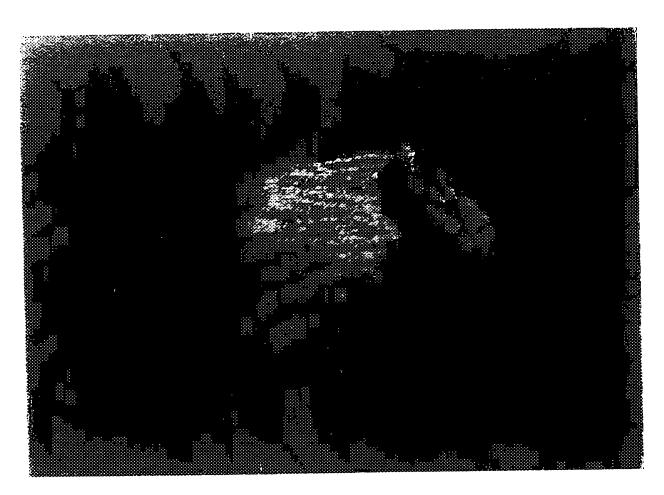
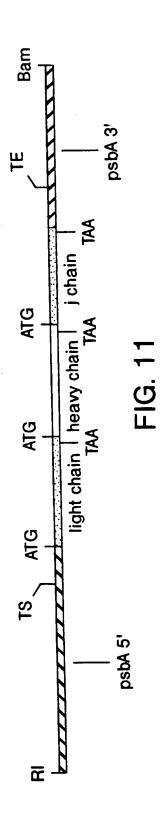


FIG. 10

an Ag

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